Jae-Hyun Yang, Ph.D.

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Research summary:

My research goals are to understand the mechanisms of aging and reverse them in order to treat age-related diseases. I am passionate about creating a network of researchers collaborating at the nexus of aging to address the growing number of elderly people suffering from chronic diseases. We have demonstrated that a loss of epigenetic information causes molecular and physiological aging in mice. Following this, reprogramming to restore youthful epigenetic information can reverse biological age. Thus, by manipulating the epigenome, aging can be driven forward and backward. Using mouse models, aging-inducible human organoids, microfluidic organ-chips, and high-throughput aging screening platforms, I aim to study rejuvenation mechanisms and develop safe age reversal therapies that restore youthful epigenetic information and tissue function.

Keywords: Mechanisms of reverse aging; Epigenetic reprogramming; Organoids; Stem cells

Education:

08/2011	Ph.D., School of Pharmacy, Sungkyunkwan University, Suwon, South Korea (Mentor: Eun-Jung Cho, Ph.D.)
02/2007	M.S., School of Pharmacy, Sungkyunkwan University, Suwon, South Korea (Mentor: Eun-Jung Cho, Ph.D.)
02/2005	B.S., Department of Biotechnology, Paichai University, South Korea

Professional experience:

12/2012-	Postdoctoral fellow, Harvard Medical School, Boston, MA, USA (Mentor: David A. Sinclair, Ph.D.)
2011-2012	Postdoctoral fellow , Sungkyunkwan University, Suwon, South Korea (Mentor: Eun-Jung Cho, Ph.D.)
2008-2009	Teaching Assistant, Sungkyunkwan University, South Korea
2005-2006	Research Assistant, Sungkyunkwan University, South Korea

Awards and honors:

11/2022	Outstanding Research Award (Junior Research Scientist), Association of Korean Neuroscientist
12/2020	Korea American Bioscience Forum 2020 NYKB Fellowship, New York Korean Biologists
03/2018	Epigenetics Initiative Travel Grants, Department of Genetics, Harvard Medical School
04/2016	Epigenetics Initiative Travel Grants, Department of Genetics, Harvard Medical School
06/2014	KASBP-Yuhan fellowship, Korean American Society in Biotech and Pharmaceuticals
12/2012	Postdoctoral fellowship, Fostering Next-generation Researchers Program, NRF of Korea
05/2011	KRIBB Best research award, Korean Society for Biochemistry and Molecular Biology
2008-2010	Brain Korea 21 scholarship, Ministry of Education, Science and Technology
04/2007	Poster award, School of Pharmacy Research Day, Sungkyunkwan University
04/2006	Presentation award, School of Pharmacy Research Day, Sungkyunkwan University

Publications:

- 1. **JH Yang**, MV Lopez, Z Chen, N Ibrahim, X Tian, PT Griffin, DA Sinclair. An organoid model of brain aging and epigenetic drift (Manuscript in preparation)
- 2. <u>JH Yang</u>, JA Amorim, M Hayano, EK Nishimura, P Oberdoerffer, DA Sinclair. The Inducible Changes to the Epigenome (ICE) system to study DNA damage response, epigenetic alteration, and aging. (Manuscript in preparation)
- 3. **JH Yang**, C Petty, MV Lopez, N Ibrahim, S Maybury-Lewis, T Dixon-McDougall, Z Chen, PT Griffin, S Angeli, DA Sinclair. Chemically induced reprogramming to reverse cellular aging. (Manuscript in preparation)

- 4. **JH Yang**^{*#}, M Hayano^{*}, TP Griffin, JA Amorim, MS Bonkowski, JK Apostolides, EL Salfati, M Blanchette, EM Munding, M Bhakta, YC Chew, W Guo, X Yang, S Maybury-Lewis, X Tian, JM Ross, G Coppotelli, Meer MV, R Rogers-Hammond, Y Lu, JW Pippin, ML Creswell, Z Dou, C Xu, DL Vera, SJ Mitchell, A Das, BL O'Connell, S Thakur, A Kane, Q Su, Y Mohri, EK Nishimura, L Schaevitz, N Garg, A Balta, MA Rego, Gregory-Ksander M, Jakobs TC, Zhong L, Wakimoto H, Andari JE, Grimm D, Mostoslavsky R, Wagers AJ, K Tsubota, SJ Bonasera, C Palmeira, JG Seidman, CE Seidman, NS Wolf, JA Kreiling, JM Sedivy, GF Murphy, RE Green, BA Garcia, SL Berger, P Oberdoerffer, SJ Shankland, VN Gladyshev, BR Ksander, AR Pfenning, LA Rajman, DA Sinclair[#]. Loss of epigenetic information as a cause of mammalian aging. *Cell* 2023 Jan 19;186(2):305-326
 - [JH Yang: Co-first & co-corresponding author, 2021 IF: 66.85]
 - Research highlights:
 - Nature aging, Loss of epigenetic information drives aging
 - Cell Metabolism, Putting aging on ICE
 - Science, Two research teams reverse signs of aging in mice
 - Signal Transduction and Targeted Therapy, The loss of epigenetic information: not only consequences but a cause of mammalian aging
 - TIME, Scientists Have Reached a Key Milestone in Learning How to Reverse Aging
 - CNN, Old mice grow young again in study. Can people do the same?
 - THE SCIENTIST, Epigenetic Manipulations Can Accelerate or Reverse Aging in Mice
 - Harvard Medical School, Loss of Epigenetic Information Can Drive Aging, Restoration Can Reverse It
 - The Harvard Gazette, Has first person to live to be 150 been born?
 - Boston Globe, New research points to a way to reverse aging. But don't expect a miracle drug overnight
 - Yale Scientific Magazine, Turning Back the Clock
 - Chosun Biz, [사이언스카페] 하버드대 한인 과학자, 회춘의 새로운 길 찾았다
- 5. HJ Lim, YH Baek, MY Park, **JH Yang**, MJ Kim, N Sung, YH Sohn, SH Lee, JE Park, YJ Yang. Performance Analysis of Self-Collected Nasal and Oral Swabs for Detection of SARS-CoV-2. *Diagnostics* 2022 Sep 21;12(10):2279 [Co-author, 2021 IF: 3.992]
- 6. YH Baek, MY Park, HJ Lim, HS Jung, **JH Yang**, YH Sohn, SH Lee, JE Park, YJ Yang. Evaluation of Alternative Transport Media for RT-qPCR-Based SARS-CoV-2 Testing. *Int J Anal Chem* 2022 Aug 10;2022:5020255 [Co-author, 2021 IF: 1.698]
- 7. HJ Lim, HS Jung, MY Park, YH Baek, B Kannappan, JY Park, **JH Yang**, JH Seol, MH Lee, SK Jung, SH Lee, JE Park, YJ Yang. Evaluation of Three Automated Extraction Systems for the Detection of SARS-CoV-2 from Clinical Respiratory Specimens. *Life* 2022 Jan 4;12(1):68 [Co-author, 2021 IF: 3.251]
- 8. HJ Lim, ER Kang, MY Park, BK Kim, MJ Kim, S Jung, KH Roh, N Sung, **JH Yang**, MW Lee, SH Lee, YJ Yang. Development of a multiplex real-time PCR assay for the simultaneous detection of four bacterial pathogens causing pneumonia. *PLoS One* 2021 Jun 17;16(6):e0253402 [Co-author, 2021 IF: 3.752]
- HJ Lim, JE Park, MY Park, JH Baek, S Jung, N Sung, JH Yang, MW Lee, SH Lee, YJ Yang. Assay System for Simultaneous Detection of SARS-CoV-2 and Other Respiratory Viruses. *Diagnostics* 2021 Jun 13;11(6):1084 [Coauthor, 2021 IF: 3.992]
- 10. Y Lu, B Brommer, X Tian, A Krishnan, M Meer, C Wang, DL Vera, Q Zeng, D Yu, MS Bonkowski, JH Yang, S Zhou, EM Hoffmann, MM Karg, MB Schultz, AE Kane, N Davidsohn, E Korobkina, K Chwalek, LA Rajman, GM Church, K Hochedlinger, VN Gladyshev, S Horvath, ME Levine, MS Gregory-Ksander, BR Ksander, Z He, DA Sinclair. Reprogramming to recover youthful epigenetic information and restore vision. *Nature* 2020 Dec 2;588(7836):124-129 [Co-author, 2020 IF: 49.962]
- J Park, H Lee, N Han, S Kwak, HT Lee, JH Kim, K Kang, BH Youn, JH Yang, HJ Jeong, JS Kang, SY Kim, JW Han, HD Youn, EJ Cho. Long noncoding RNA ChRO1 facilitates DAXX-dependent H3.3 deposition for transcriptionassociated heterochromatin reorganization. *Nucleic Acids Res* 2018 Dec 14;46(22):11759-11775 [Co-author, 2018 IF: 11.147]
- 12. <u>JH Yang</u>^{*}, TY Song^{*}, C Jo^{*}, J Park, HY Lee, I Song, S Hong, KY Jung, J Kim, HD Youn, JW Han, EJ Cho. Differential regulation of the histone chaperone HIRA during muscle cell differentiation by a phosphorylation switch. *Exp Mol Med* 2016 Aug 12;48(8):e252 [JH Yang: Co-first author, 2016 IF: 5.063]
- 13. JH Seol, TY Song, SE Oh, C Jo, A Choi, B Kim, J Park, S Hong, I Song, KY Jung, **JH Yang**, H Park, JH Ahn, JW Han, EJ Cho. Identification of small molecules that inhibit the histone chaperone Asf1 and its chromatin function. *BMB Rep* 2015 Dec;48(12):685-690 [Co-author, 2015 IF: 2.782]
- 14. Y Song, JH Seol, **JH Yang**, HJ Kim, JW Han, HD Youn, EJ Cho. Dissecting the roles of the histone chaperones reveals the evolutionary conserved mechanism of transcription-coupled deposition of H3.3. *Nucleic Acids Res* 2013 May 1;41(10):5199-209 [Co-author, 2013 IF: 9.112]

- 15. YJ Yang, TY Song, J Park, J Lee, J Lim, H Jang, YN Kim, **JH Yang**, Y Song, A Choi, HY Lee, CH Jo, JW Han, ST Kim, HD Youn, EJ Cho. Menin mediates epigenetic regulation via histone H3 lysine 9 methylation. *Cell Death Dis* 2013 Apr 11;4(4):e583 [Co-author, 2013 IF: 5.177]
- 16. TY Song, **JH Yang**, JY Park, Y Song, JW Han, HD Youn, EJ Cho. The role of histone chaperones in osteoblastic differentiation of C2C12 myoblasts. *Biochem Biophys Res Commun* 2012 Jul 13;423(4):726-732 [Co-author, 2013 IF: 2.281]
- 17. JH Yang, JH Choi, H Jang, JY Park, JW Han, HD Youn, EJ Cho. Histone chaperones cooperate to mediate Mef2targeted transcriptional regulation during skeletal myogenesis. *Biochem Biophys Res Commun* 2011 Apr 15;407(3):541-547 [JH Yang: First author, 2011 IF: 2.484]
- JH Yang, Y Song, JH Seol, JY Park, YJ Yang, JW Han, HD Youn, EJ Cho. Myogenic transcriptional activation of *MyoD* mediated by replication-independent histone deposition. *Proc Natl Acad Sci U S A* 2011 Jan 4;108(1):85-90 [JH Yang: First author, 2011 IF: 9.681]

Talks:

05/2023	Evidence for forward and reverse aging via epigenetic manipulation Department of Biomedical Engineering Colloquium, UNIST, Ulsan, Korea
05/2022	Evidence for forward and reverse aging via epigenetic manipulation TEU (TIDE Envision University)-MED 3, (virtual lecture)
03/2023	Loss of epigenetic information as a cause of mammalian aging Stochastic Stemness Research Center, College of Medicine, Seoul National University (virtual)
03/2023	Loss of epigenetic information as a cause of mammalian aging College of Medicine, Korea University (virtual)
02/2023	Loss of epigenetic information as a cause of mammalian aging 300th New England Bioscience Society (NEBS) monthly meeting, Boston, MA, USA
02/2023	Loss of epigenetic information as a cause of mammalian aging School of Biological Sciences, Seoul National University (virtual)
02/2023	Loss of epigenetic information as a cause of mammalian aging Department of Microbiology & Immunology, College of Medicine, Seoul National University (virtual)
01/2023	Loss of epigenetic information as a cause of mammalian aging College of Medicine, Yonsei University (virtual)
09/2022	Evidence for epigenetic drift as a reversible cause of aging The 31st International KOGO Annual Conference (virtual)
09/2022	Evidence for epigenetic drift as a reversible cause of aging Data & Journal Club, Department of Genetics, Harvard Medical School, Boston, MA, USA
06/2022	The epigenetics of aging — how to reset the aging clock 2022 UST Global Mentoring Conference, (virtual mentoring)
05/2022	The epigenetics of aging TEU (TIDE Envision University)-MED 2, (virtual lecture)
02/2022	Evidence for epigenetic drift as a reversible cause of aging 2022 SKKU-GSBMS International Symposium(Metabolism, Aging, Mitochondria, and AI), (virtual)
11/2021	The epigenetics of aging: why and how we age TEU (TIDE Envision University)-MED 1, (virtual lecture)
08/2021	Loss of epigenetic information as a cause of mammalian aging NEBS-Seoul National University College of Medicine joint symposium (virtual)
07/2021	Loss of epigenetic information as a cause of mammalian aging NEBS-Yonsei University College of Medicine joint seminar (virtual)
07/2021	Faithful DNA repair causes the erosion of the epigenetic landscape and aging NEBS-Seoul National University College of Pharmacy joint symposium (virtual)
12/2020	Loss of epigenetic information as a cause of mammalian aging KOREAN-AMERICAN BIOSCIENCE FORUM 2020 (virtual)

09/2020	DNA break-induced loss of epigenetic information as a cause of mammalian aging Mechanisms of Aging - Cold Spring Harbor Laboratory (virtual)
11/2019	DNA break-induced epigenetic drift as a cause of mammalian aging Data & Journal Club, Department of Genetics, Harvard Medical School, Boston, MA, USA
11/2019	Erosion of the epigenetic landscape and loss of cellular identity as a cause of aging in mammals Fifth Annual Northeastern Glenn Symposium on the Biology of Aging, Farmington, CT, USA
10/2019	DNA break-induced epigenetic drift as a cause of mammalian aging ASHG 2019 (American Society of Human Genetics) CoLab, Houston, Tx, USA
08/2019	Loss of epigenetic information as a cause of mammalian aging Korea Institute of Oriental Medicine, Daegu, South Korea
08/2019	Loss of epigenetic information as a cause of mammalian aging Graduate School of Medical Science and Engineering, KAIST, Daejeon, South Korea
08/2019	Loss of epigenetic information as a cause of mammalian aging Department of Biomedical Engineering, School of Life Sciences, UNIST, Ulsan, South Korea
08/2019	Loss of epigenetic information as a cause of mammalian aging School of Pharmacy, Sungkyunkwan University, Suwon, South Korea
08/2019	Loss of epigenetic information as a cause of mammalian aging School of Medicine, Kyungpook National University, Daegu, South Korea
08/2019	Loss of epigenetic information as a cause of mammalian aging College of Medicine, Ajou University, Suwon, South Korea
09/2017	Evidence for epigenetic change as a cause of mammalian aging Data & Journal Club, Department of Genetics, Harvard Medical School, Boston, MA, USA
05/2016	Evidence for epigenomic change as a cause of mammalian aging Keystone Symposia, Epigenetic and Metabolic Regulation of Aging (E1), Santa Fe, NM, USA
08/2015	Evidence for histone depletion as a cause of mammalian aging Data & Journal Club, Department of Genetics, Harvard Medical School, Boston, MA, USA
01/2011	Myogenic transcriptional activation of <i>MyoD</i> mediated by RI histone deposition Chromatin & Epigenetics symposium, KSBMB, Hongcheon-gun, South Korea
01/2009	Histone chaperone HIRA regulates <i>MyoD</i> during muscle differentiation Chromatin & Epigenetics symposium, KSBMB, Hongcheon-gun, South Korea
11/2006	Transcriptional Regulation by Histone Chaperone HIRA during Myogenesis School of Pharmacy Research Day, Sungkyunkwan University, Suwon, South Korea
Posters:	
11/2022	DNA break-induced loss of epigenetic information as a cause of neuronal aging Society for Neuroscience (SfN 2022), San Diego, CA, USA
09/2022	Evidence for loss of epigenetic information as a reversible cause of aging in mammals Mechanisms of Aging - Cold Spring Harbor Laboratory, Cold Spring Harbor, NY, USA
10/2020	DNA break-induced epigenetic drift as a cause of neuronal aging Neuroepigenetics: From Cells to Behaviour and Disease-EMBO (virtual)
10/2020	Loss of epigenetic information as a cause of mammalian aging 2020 KASBP Fall eSymposium (virtual)
05/2020	DNA break-induced loss of epigenetic information leads to aging in mammals Genome Organization & Nuclear Function-Cold Spring Harbor Laboratory (virtual)
11/2019	DNA Break-Induced Epigenetic Drift as a Cause of Mammalian Aging Korean American Society in Biotech and Pharmaceuticals Fall Symposium, Andover, MA, USA
11/2018	Evidence that epigenetic alterations drive mammalian aging GSA's 2018 Annual Scientific Meeting, Boston, MA, USA
09/2018	Evidence for epigenetic noise and loss of cell identity as a cause of aging Mechanisms of Aging - Cold Spring Harbor Laboratory, Cold Spring Harbor, NY, USA

07/2018	Evidence for epigenetic change as a cause of mammalian aging Gordon Research Conference on Chromatin Structure and Function, Newry, ME, USA
09/2016	Evidence for epigenomic change as a cause of mammalian aging Mechanisms of Aging - Cold Spring Harbor Laboratory, Cold Spring Harbor, NY, USA
05/2016	Evidence for epigenomic change as a cause of mammalian aging Keystone Symposia, Epigenetic and Metabolic Regulation of Aging (E1), Santa Fe, NM, USA
12/2014	Evidence That The Cellular Response to Broken DNA Drives the Chromatin and Gene Expression Changes during Aging HMS Epigenetics Symposium, Boston, MA, USA
09/2014	Evidence that the cellular response to broken DNA drives the chromatin and gene expression changes during aging Molecular Genetics of Aging - Cold Spring Harbor Laboratory, Cold Spring Harbor, NY, USA
07/2014	A new driver of aging: the reorganization of chromatin structure Korean American Society in Biotech and Pharmaceuticals Spring Symposium, Edison, NJ, USA
06/2012	The role of HIRA phosphorylation in H3.3 incorporation The 12 th Asian Conference on Transcription, Jeju, South Korea
05/2011	Histone chaperones cooperate to mediate Mef2-targeted transcriptional regulation during skeletal myogenesis The Korean Society for Biochemistry and Molecular Biology, Seoul, South Korea
05/2010	Myogenic activation of <i>MyoD</i> with variant histone H3.3 CSH Asia Conference: Epigenetics, Chromatin & Transcription, Suzhou, China
10/2009	Histone chaperone HIRA regulates MyoD expression during muscle differentiation The Korean Society for Molecular and Cellular Biology, Seoul, South Korea
10/2008	Myogenic regulation of muscle specific genes by histone chaperones ASBMB, Transcriptional Regulation by Chromatin and RNA Polymerase II, Tahoe City, CA, USA
10/2006	Myogenic regulation by histone chaperone HIRA The Korean Society of Medical Biochemistry and Molecular Biology, Seoul, South Korea

Research projects:

2012-2022	Role of epigenetic decay in cell senescence and aging (5R01AG019719), National Institute on Aging, \$2,972,204, Co-investigator
2012-2023	Research fund (06775092-01), Glenn Foundation for Medical Research, \$2,000,000, Co-investigator
2012-2021	SIRT1 as a regulator of health and lifespan of mammals (5R37AG028730), National Institute on Aging, \$4,128,877, Co-investigator

Editorial activities:

11/2022-	Editorial board (Review editor), Frontiers in Aging
02/2022	Co-reviewer, Nature
05/2021	Co-reviewer, Cell Reports
04/2020	Co-reviewer, Trends in Genetics
03/2018	Co-reviewer, Cell Metabolism
09/2016	Co-reviewer, Molecular Cell
08/2016	Co-reviewer, Molecular Cell
04/2016	Co-reviewer, Science Advances
03/2016	Co-reviewer, Molecular Cell
02/2016	Co-reviewer, <i>eLife</i>
02/2015	Co-reviewer, BMC Genomics
10/2013	Co-reviewer, Current Biology

Professional memberships:

2022-	Society for Neuroscience
2019-2021	Korean American Society in Biotech and Pharmaceuticals
2018-2019	The Gerontological Society of America
2014-2015	Korean American Society in Biotech and Pharmaceuticals
2009-2012	Korean Society for Biochemistry and Molecular Biology
2008-2012	Korean Society for Molecular and Cellular Biology
2007-2012	The Pharmaceutical Society of Korea

Workshops and courses:

06-08/2018Funding Your Research: NIH, Harvard Catalyst Postgraduate Education08/2018Pathway Analysis of Genomic data using MetaCore, Harvard Medical School06/2018Introduction to ChIP-Seq and data analysis using Galaxy, Harvard Medical School05/2018Genome Engineering 2018, Broad Institute, Boston, MA, USA04-05/2018Introduction to R, Harvard Chan Bioinformatics Core07/2017Imaging Solutions for Scientific Communication, Harvard Medical School09/2015Intro R/Bioconductor, Harvard Medical School08/2015Bioinformatics summer course Cell Profiler, Harvard Medical School05/2015Consume Tenging Scientific Q. Dure d Medical School
06/2018Introduction to ChIP-Seq and data analysis using Galaxy, Harvard Medical School05/2018Genome Engineering 2018, Broad Institute, Boston, MA, USA04-05/2018Introduction to R, Harvard Chan Bioinformatics Core07/2017Imaging Solutions for Scientific Communication, Harvard Medical School09/2015Intro to R/Bioconductor, Harvard Medical School08/2015Bioinformatics summer course Cell Profiler, Harvard Medical School
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05/2015 Compared Euclidean 2.0 Due of Luctitude
05/2015 Genome Engineering 3.0, Broad Institute
05/2015 MetaCore "omics" analysis, HMS/HSDM Office for Postdoctoral Fellows
03/2015 ChIP-seq analysis, Harvard School of Public Health Bioinformatics Core
11/2014 NGS Data: Standard data processing and workflow analysis, Harvard Medical School
05/2014 HMS ChIP-Seq Workshop, HMS/HSDM Office for Postdoctoral Fellows
04/2014 mRNA-seq analysis using JMP Genomics software, Harvard Medical School

Mentorship:

01/2023-	Yashi Singh (Northeastern University), co-op student in the Sinclair lab
08/2022-04/2023	Nabilah Ibrahim (Northeastern University), co-op student in the Sinclair lab
07/2022-04/2023	Zhili Chen (Zhejiang University), student intern in the Sinclair lab
06-08/2022	Sally Tabakh (Noble and Greenough School), HMS Genetics DEI Internship Program
01-08/2022	Maria Vina Lopez (University of Maine), student intern in the Sinclair lab
10-12/2019	Kejun Ying (Harvard University), rotation student in the Sinclair lab
04-06/2018	Alex Plesa (Harvard Medical School), rotation student in the Sinclair lab
01-12/2018	Patrick Griffin (Harvard Medical School), graduate student in the Sinclair lab
01-12/2017	Yuancheng Lu (Harvard Medical School), graduate student in the Sinclair lab
05-08/2017	Phillip Dmitriev (University of Florida), summer student in the Sinclair lab
01-02/2016	Andrew Kane (Harvard University), rotation student in the Sinclair lab
05-08/2015	Sarah Hemphill (Middlebury College), summer student in the Sinclair lab

References:

David A. Sinclair, A.O., Ph.D.	Co-Director, Paul F. Glenn Center for Biology of Aging Research Professor, Department of Genetics – Blavatnik Institute, Harvard Medical School 77 Avenue Louis Pasteur, NRB-931B, Boston, MA 02115, USA Email: <u>david_sinclair@hms.harvard.edu</u>
Eun-Jung Cho, Ph.D.	Professor, School of Pharmacy, Sungkyunkwan University Seoburo 2066, Jangan-gu, Suwon, Gyeonggi-do 16419, Republic of Korea Email: <u>echo@skku.edu</u>
Vadim N. Gladyshev, Ph.D.	Professor of Medicine, Harvard Medical School Director, Center for Redox Medicine, Brigham And Women's Hospital 77 Avenue Louis Pasteur, NRB-435, Boston, MA 02115, USA Email: <u>vgladyshev@rics.bwh.harvard.edu</u>